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L10: Entry 2 of 2

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May 1, 2003

DERWENT-ACC-NO: 2002-373930

DERWENT-WEEK: 200331

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TITLE: Identifying agents for treatment or prevention of cytomegalovirus infection, comprises contacting test compound with cellular kinase and detecting change in cellular kinase activity

INVENTOR: BEVEC, D; HABENBERGER, P ; SCHUBART, D ; STEIN-GERLACH, M

PRIORITY-DATA: 2000US-240750P (October 16, 2000), 2001US-0981397 (October 16, 2001)

 [Search Selected](#) [Search ALL](#) [Clear](#)**PATENT-FAMILY:**

PUB-NO	PUB-DATE	LANGUAGE	PAGES	MAIN-IPC
<input type="checkbox"/> US 20030082519 A1	May 1, 2003		000	A61K039/395
<input type="checkbox"/> EP 1201765 A2	May 2, 2002	E	049	C12Q001/48

INT-CL (IPC): A01 N 61/00; A61 K 31/00; A61 K 39/395; A61 K 48/00; A61 P 31/12; C07 K 16/00; C12 P 21/06; C12 Q 1/48; C12 Q 1/68; C12 Q 1/70; G01 N 33/53

ABSTRACTED-PUB-NO: EP 1201765A

BASIC-ABSTRACT:

NOVELTY - Identifying compounds (A) for treating and/or preventing cytomegalovirus (CMV) infection and/or related diseases comprising contacting a test compound with at least one of the cellular kinases RICK, RIP, Nck-Interacting kinase, MKK3 and SRPK-2 (undefined) and detecting any change in kinase activity, is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

- (1) detecting CMV infection and/or related diseases by detecting activity of any of the specified kinases in a patient sample, cells or cell lysates;
- (2) mono- or poly-clonal antibodies (Ab) that bind to any of the specified kinases;
- (3) preventing and/or treating CMV infection or related diseases, or for regulating production of CMV in an individual or cells, by administering an inhibitor or activator of any of the specified kinases;
- (4) oligonucleotides (ON) that bind to RNA or DNA encoding any of the specified kinases;
- (5) regulating expression of any of the specified kinases by administering to an

individual, or cells, an inhibitor or activator of transcription of the relevant DNA or translation of the relevant RNA;

(6) solid support for detecting CMV infection in an individual or cell comprising at least one immobilized ON able to detect activity of any of the specified kinases; and

(7) solid support for screening (A) comprising one or more immobilized ON that encode any of the specified kinases or these kinases themselves.

ACTIVITY - Virucide. RICK was transiently overexpressed, as a fusion with a hemagglutinin (HA) tag, in human embryonic kidney 293 cells, then immunoprecipitated (anti-HA antibody and protein A-Sepharose). The beads were washed, then tested for kinase activity by incubation in a mixture containing gamma (33P)-adenosine triphosphate and various concentrations of 8-methyl-6-phenyl-2-(pyridin-4-ylamino)-8H-pyrido(2,3-d)pyrimidin-7-one (Aa). After 30 minutes at 30 deg. C, reaction was stopped and phosphorylation determined by electrophoresis and autoradiography. (Aa) has an inhibitory concentration (IC)50 for inhibition of RICK of 500 nM and for inhibition of CMV of 1.4 micro M.

MECHANISM OF ACTION - Modulation of cellular kinases that are specifically upregulated during CMV infection.

USE - (A) are used to treat and/or prevent CMV infections and related diseases. Oligonucleotides that can detect the specified kinases can also be used for diagnosis of infection.

ABSTRACTED-PUB-NO: EP 1201765A

EQUIVALENT-ABSTRACTS:

CHOSEN-DRAWING: Dwg.0/2

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L10 schubart d.in. 2

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L9 schubart d.in. 0

L8 kinase RIP and CMV.clm. 1

L7 Rip kinase and CMV.clm. 0

L6 Rip kinases and CMV.clm. 0

L5 kinases and CMV.clm. 217

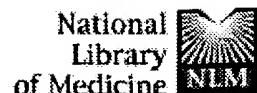
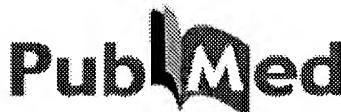
L4 kinases and CMV 5380

L3 L1 and virus 1

L2 L1 and cmv 0

L1 6558903.bn. 1

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<input type="checkbox"/> 1: Yang Y, Ma J, Chen Y, Wu M.	Related Articles , Links
Nucleocytoplasmic shuttling of RIP3: Identification of novel nuclear export and import signals in RIP3. J Biol Chem. 2004 Jun 18 [Epub ahead of print] PMID: 15208320 [PubMed - as supplied by publisher]	
<input type="checkbox"/> 2: Hu WH, Mo XM, Walters WM, Brambilla R, Bethea JR.	Related Articles , Links
TNAP, a novel repressor of NF-kappa B-inducing kinase, suppresses NF-kappa B activation. J Biol Chem. 2004 Jun 18 [Epub ahead of print] PMID: 15208311 [PubMed - as supplied by publisher]	
<input type="checkbox"/> 3: Shen HM, Lin Y, Choksi S, Tran J, Jin T, Chang L, Karin M, Zhang J, Liu ZG.	Related Articles , Links
Essential roles of receptor-interacting protein and TRAF2 in oxidative stress-induced cell death. Mol Cell Biol. 2004 Jul;24(13):5914-22. PMID: 15199146 [PubMed - in process]	
<input type="checkbox"/> 4: Zha J, Zhou Q, Xu LG, Chen D, Li L, Zhai Z, Shu HB.	Related Articles , Links
RIP5 is a RIP-homologous inducer of cell death. Biochem Biophys Res Commun. 2004 Jun 25;319(2):298-303. PMID: 15178406 [PubMed - in process]	
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The kinase activity of Rip 1 is not required for TNF-alpha-induced Ikk or p38 MAP kinase activation or for the ubiquitination of Rip 1 by Traf 2. J Biol Chem. 2004 Jun 1 [Epub ahead of print] PMID: 15175328 [PubMed - as supplied by publisher]	
<input type="checkbox"/> 6: Kurenova E, Xu LH, Yang X, Baldwin AS Jr, Craven RJ, Hanks SK, Liu ZG, Cance WG.	Related Articles , Links
Focal adhesion kinase suppresses apoptosis by binding to the death domain of receptor-interacting protein. Mol Cell Biol. 2004 May;24(10):4361-71. PMID: 15121855 [PubMed - indexed for MEDLINE]	
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Selective Regulation by $\{\delta\}$ -PKC and PI 3-kinase in the Assembly of the Anti-Apoptotic TNFR-1 Signaling Complex in Neutrophils. Am J Physiol Cell Physiol. 2004 Apr 28 [Epub ahead of print] PMID: 15115707 [PubMed - as supplied by publisher]	

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RIP1 is an essential mediator of Toll-like receptor 3-induced NF-kappa B activation.
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PMID: 15064760 [PubMed - indexed for MEDLINE]

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Ligand-regulated association of ErbB-4 to the transcriptional co-activator YAP65 controls transcription at the nuclear level.
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Regulation of apoptosis proteins in cancer cells by ubiquitin.
Oncogene. 2004 Mar 15;23(11):2009-15. Review.
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Caspase involvement in RIP-associated CD95-induced T cell apoptosis.
Cell Immunol. 2003 Dec;226(2):78-85.
PMID: 14962495 [PubMed - indexed for MEDLINE]

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ZNF216 Is an A20-like and IkappaB kinase gamma-interacting inhibitor of NFkappaB activation.
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Mechanisms of the TRIF-induced interferon-stimulated response element

=> d 18 1-2

L8 ANSWER 1 OF 2 USPATFULL on STN
AN 2004:63787 USPATFULL
TI 14171 Protein kinase, a novel human protein kinase and uses thereof
IN Kapeller-Libermann, Rosana, Chestnut Hill, MA, UNITED STATES
PA Millennium Pharmaceuticals, Inc. (U.S. corporation)
PI US 2004048305 A1 20040311
AI US 2003-658904. A1 20030910 (10)
RLI Continuation-in-part of Ser. No. US 2001-781882, filed on 12 Feb 2001,
GRANTED, Pat. No. US 6630335
PRAI US 2000-182096P 20000211 (60)
DT Utility
FS APPLICATION
LN.CNT 5414
INCL INCLM: 435/006.000
INCLS: 435/069.100; 435/194.000; 435/320.100; 435/325.000; 536/023.200
NCL NCLM: 435/006.000
NCLS: 435/069.100; 435/194.000; 435/320.100; 435/325.000; 536/023.200
IC [7]
ICM: C12Q001-68
ICS: C07H021-04; C12N009-12
CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L8 ANSWER 2 OF 2 USPATFULL on STN
AN 2003:120036 USPATFULL
TI Cellular kinases involved in **Cytomegalovirus** infection and
their inhibition
IN Schubart, Daniel, Weil am Rhein, GERMANY, FEDERAL REPUBLIC OF
Habenberger, Peter, Munchen, GERMANY, FEDERAL REPUBLIC OF
Stein-Gerlach, Matthias, Munchen, GERMANY, FEDERAL REPUBLIC OF
Bevec, Dorian, Germerring, GERMANY, FEDERAL REPUBLIC OF
PI US 2003082519 A1 20030501
AI US 2001-981397 A1 20011016 (9)
PRAI US 2000-240750P 20001016 (60)
DT Utility
FS APPLICATION
LN.CNT 2300
INCL INCLM: 435/005.000
INCLS: 424/146.100; 514/001.000; 435/007.100; 435/069.100
NCL NCLM: 435/005.000
NCLS: 424/146.100; 514/001.000; 435/007.100; 435/069.100
IC [7]
ICM: A61K039-395
ICS: C12Q001-70; A61K031-00; A01N061-00; G01N033-53; C12P021-06
CAS INDEXING IS AVAILABLE FOR THIS PATENT.

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FILE 'MEDLINE' ENTERED AT 15:12:29 ON 06 JUL 2004

L1 3 S RIP KINASE
L2 18 S KINASE RIP
L3 26525 S CYTOMEGALOVIRUS
L4 0 S L2 AND L3
L5 0 S L1 AND L3

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SEA RIP KINASE AND CYTOMEGALOVIRUS

1 FILE CAPLUS
2 FILE USPATFULL
L6 QUE RIP KINASE AND CYTOMEGALOVIRUS

FILE 'CAPLUS' ENTERED AT 15:15:08 ON 06 JUL 2004
L7 1 S RIP KINASE AND CYTOMEGALOVIRUS

FILE 'USPATFULL' ENTERED AT 15:15:38 ON 06 JUL 2004
L8 2 S RIP KINASE AND CYTOMEGALOVIRUS